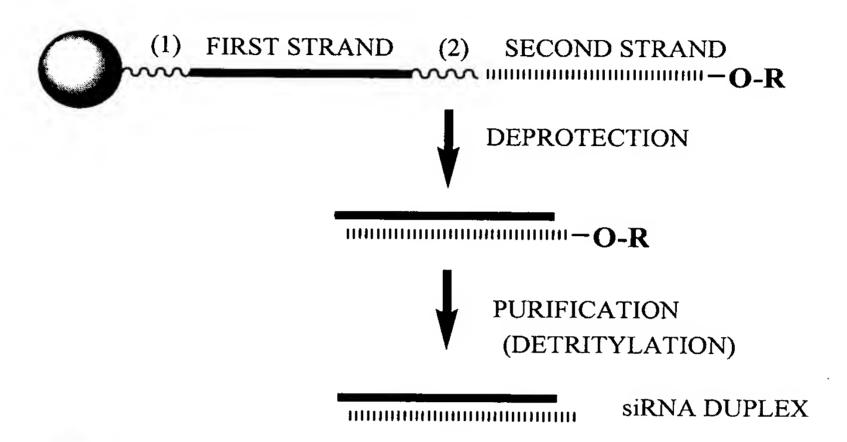
Figure 1



= SOLID SUPPORT

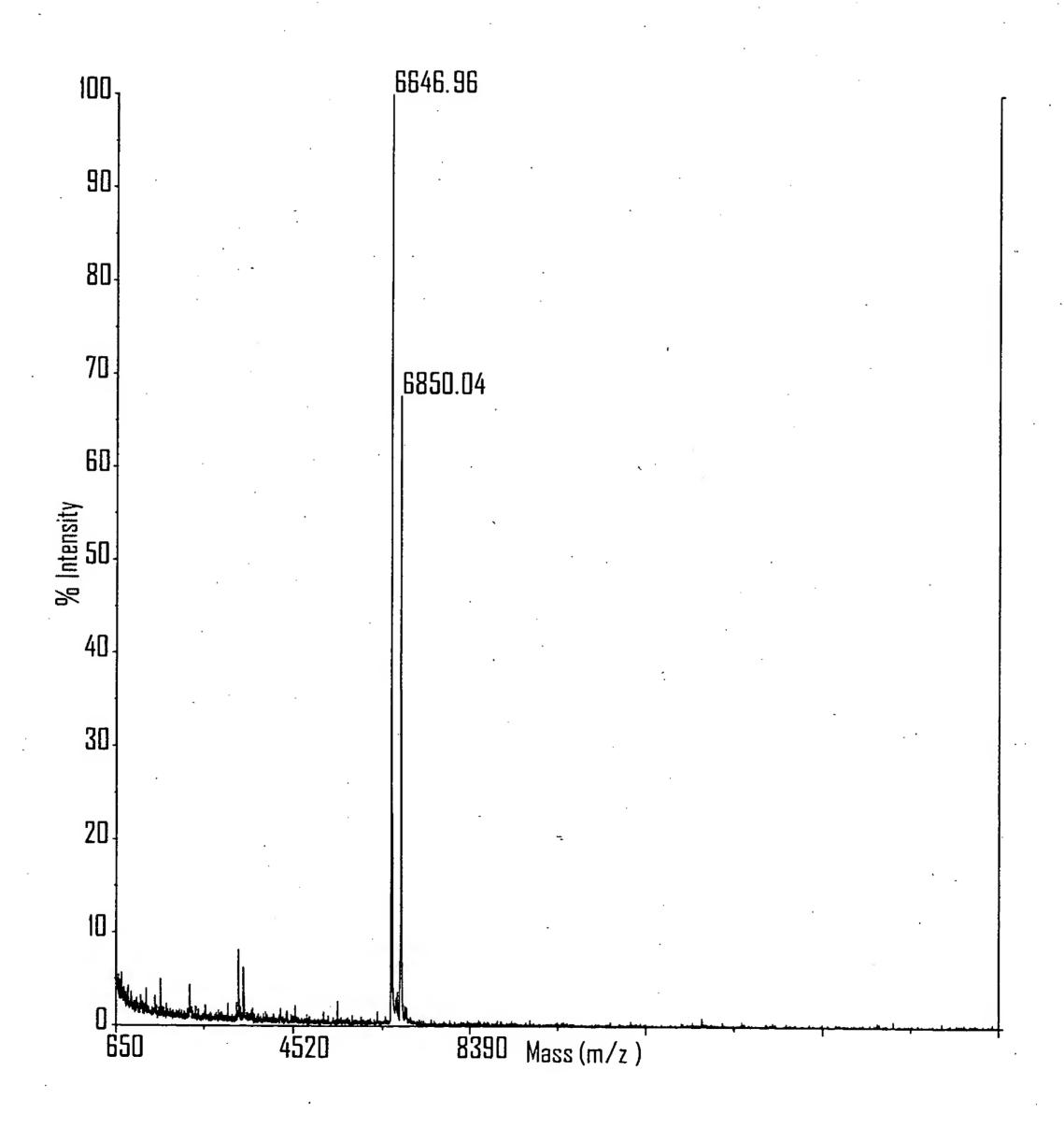
R = TERMINAL PROTECTING GROUP FOR EXAMPLE: DIMETHOXYTRITYL (DMT)

(FOR EXAMPLE: NUCLEOTIDE SUCCINATE OR INVERTED DEOXYABASIC SUCCINATE)

INVERTED DEOXYABASIC SUCCINATE LINKAGE

GLYCERYL SUCCINATE LINKAGE

Figure 2



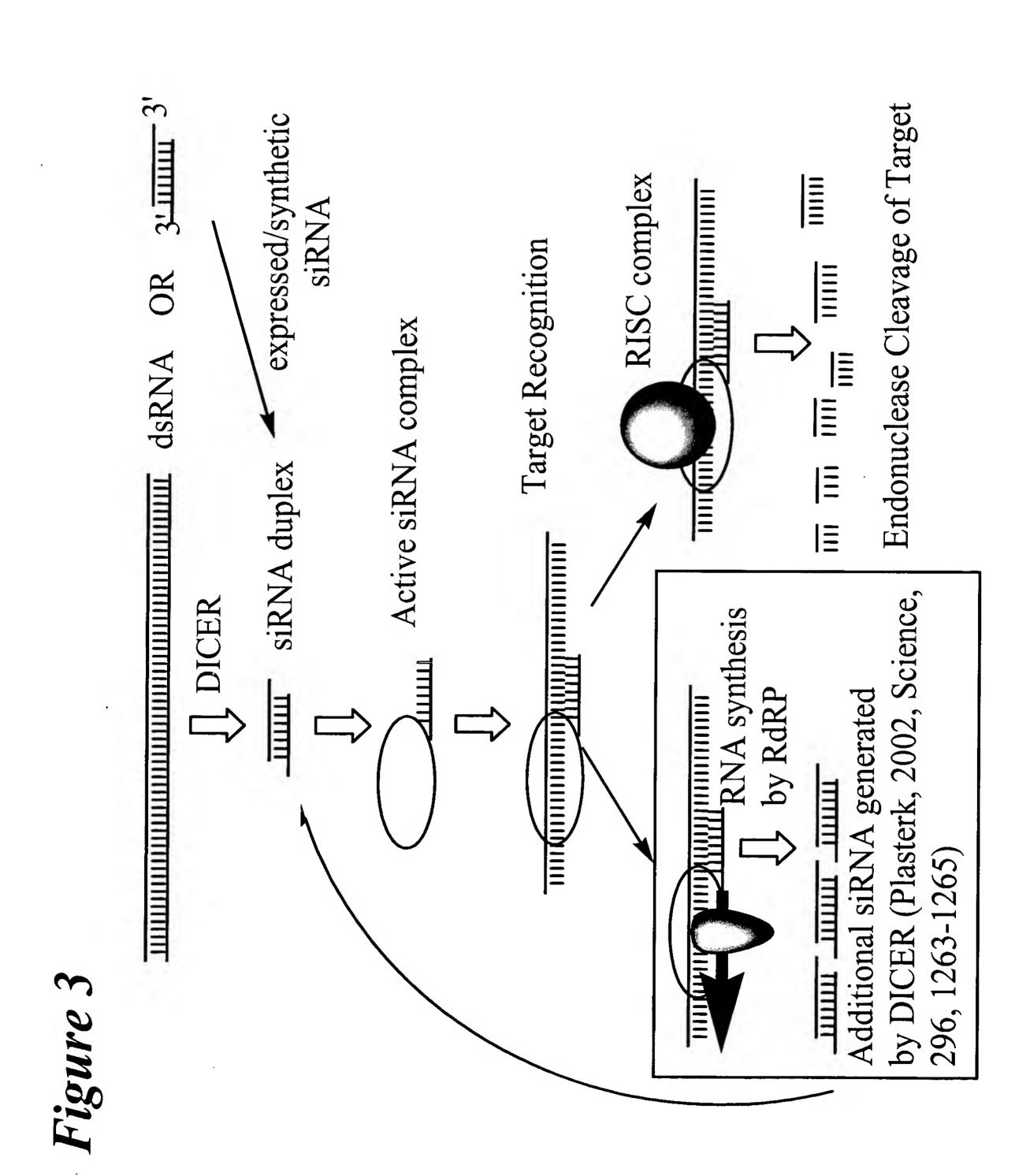


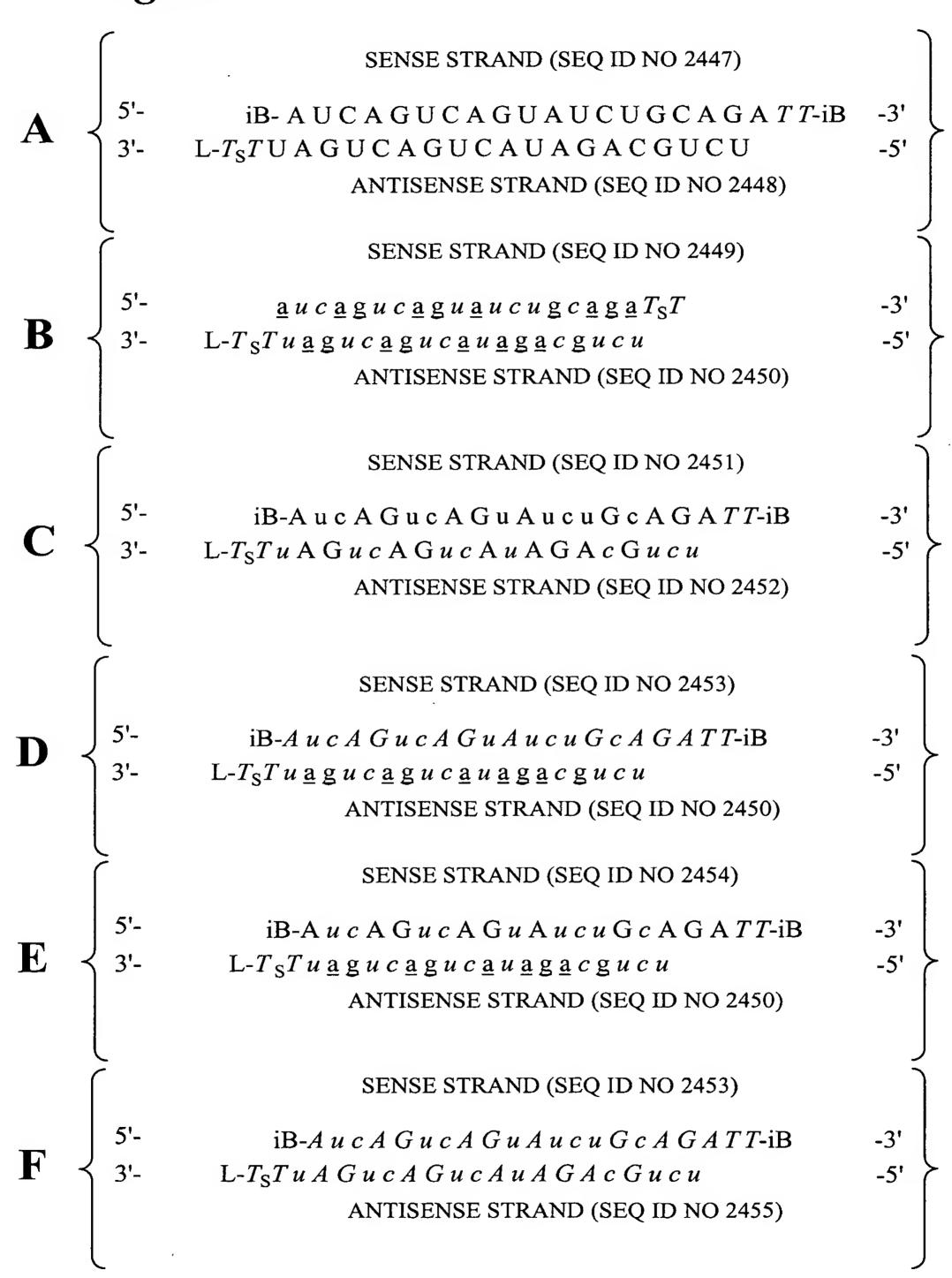
Figure 4

```
SENSE STRAND (SEQ ID NO 2438)
                ALL POSITIONS RIBONUCLEOTIDE EXCEPT POSITIONS (N N)
                B-NNNNNNNNNNNNNNNNNNNNNNN-B
           -5'
                         ANTISENSE STRAND (SEQ ID NO 2439)
                  ALL POSITIONS RIBONUCLEOTIDE EXCEPT POSITIONS (N N)
                        SENSE STRAND (SEQ ID NO 2440)
       ALL PYRIMIDINES = 2'-FLUORO AND ALL PURINES = 2'-OM EXCEPT POSITIONS (N N)
                -3'
B
            L-(N<sub>s</sub>N) NNNNNNNNNNNNNNNNNNNNN
                                                            -5'
                      ANTISENSE STRAND (SEQ ID NO 2441)
       ALL PYRIMIDINES = 2'-FLUORO AND ALL PURINES = 2'-O-ME EXCEPT POSITIONS (N N)
                         SENSE STRAND (SEQ ID NO 2442)
              ALL PYRIMIDINES = 2'-O-ME OR 2'-FLUORO EXCEPT POSITIONS (N N)
               -5'
                         ANTISENSE STRAND (SEQ ID NO 2443)
                   ALL PYRIMIDINES = 2'-FLUORO EXCEPT POSITIONS (N N)
                       SENSE STRAND (SEQ ID NO 2444)
      ALL PYRIMIDINES = 2'-FLUORO EXCEPT POSITIONS (N N) AND ALL PURINES = 2'-DEOXY
               D
           L-(N<sub>s</sub>N) NNNNNNNNNNNNNNNNNNN
                                                           -5'
                      ANTISENSE STRAND (SEQ ID NO 2441)
       ALL PYRIMIDINES = 2'-FLUORO AND ALL PURINES = 2'-O-ME EXCEPT POSITIONS (N N)
                         SENSE STRAND (SEQ ID NO 2445)
                 ALL PYRIMIDINES = 2'-FLUORO EXCEPT POSITIONS (N N)
      5'-
               B-NNNNNNNNNNNNNNNNNNNNN-B
\mathbf{E}
         -5'
                      ANTISENSE STRAND (SEQ ID NO 2441)
      ALL PYRIMIDINES = 2'-FLUORO AND ALL PURINES = 2'-O-ME EXCEPT POSITIONS (N N)
                       SENSE STRAND (SEQ ID NO 2444)
     ALL PYRIMIDINES = 2'-FLUORO EXCEPT POSITIONS (N N) AND ALL PURINES = 2'-DEOXY
              F
                                                           -3'
           -5'
                     ANTISENSE STRAND (SEQ ID NO 2446)
     ALL PYRIMIDINES = 2'-FLUORO EXCEPT POSITIONS (N N) AND ALL PURINES = 2'-DEOXY
```

POSITIONS (NN) CAN COMPRISE ANY NUCLEOTIDE, SUCH AS DEOXYNUCLEOTIDES (eg. THYMIDINE) OR UNIVERSAL BASES

- B = ABASIC, INVERTED ABASIC, INVERTED NUCLEOTIDE OR OTHER TERMINAL CAP THAT IS OPTIONALLY PRESENT
- L = GLYCERYL MOIETY THAT IS OPTIONALLY PRESENT
- S = PHOSPHOROTHIOATE OR PHOSPHORODITHIOATE

Figure 5



lower case = 2'-O-Methyl or 2'-deoxy-2'-fluoro italic lower case = 2'-deoxy-2'-fluoro underline = 2'-O-methyl

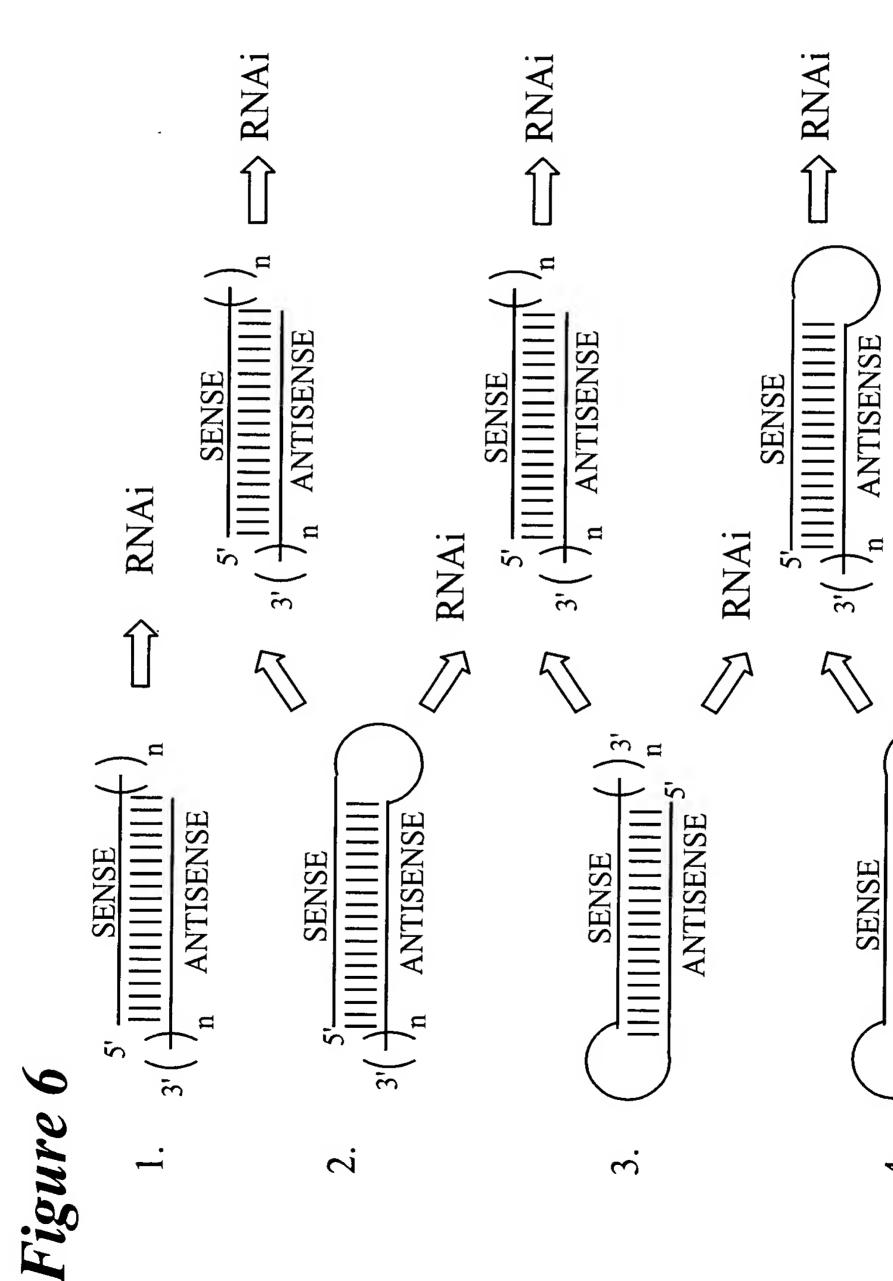
ITALIC UPPER CASE = DEOXY

B = INVERTED DEOXYABASIC

L = GLYCERYL MOIETY OPTIONALLY PRESENT

S = PHOSPHOROTHIOATE OR

PHOSPHORODITHIOATE

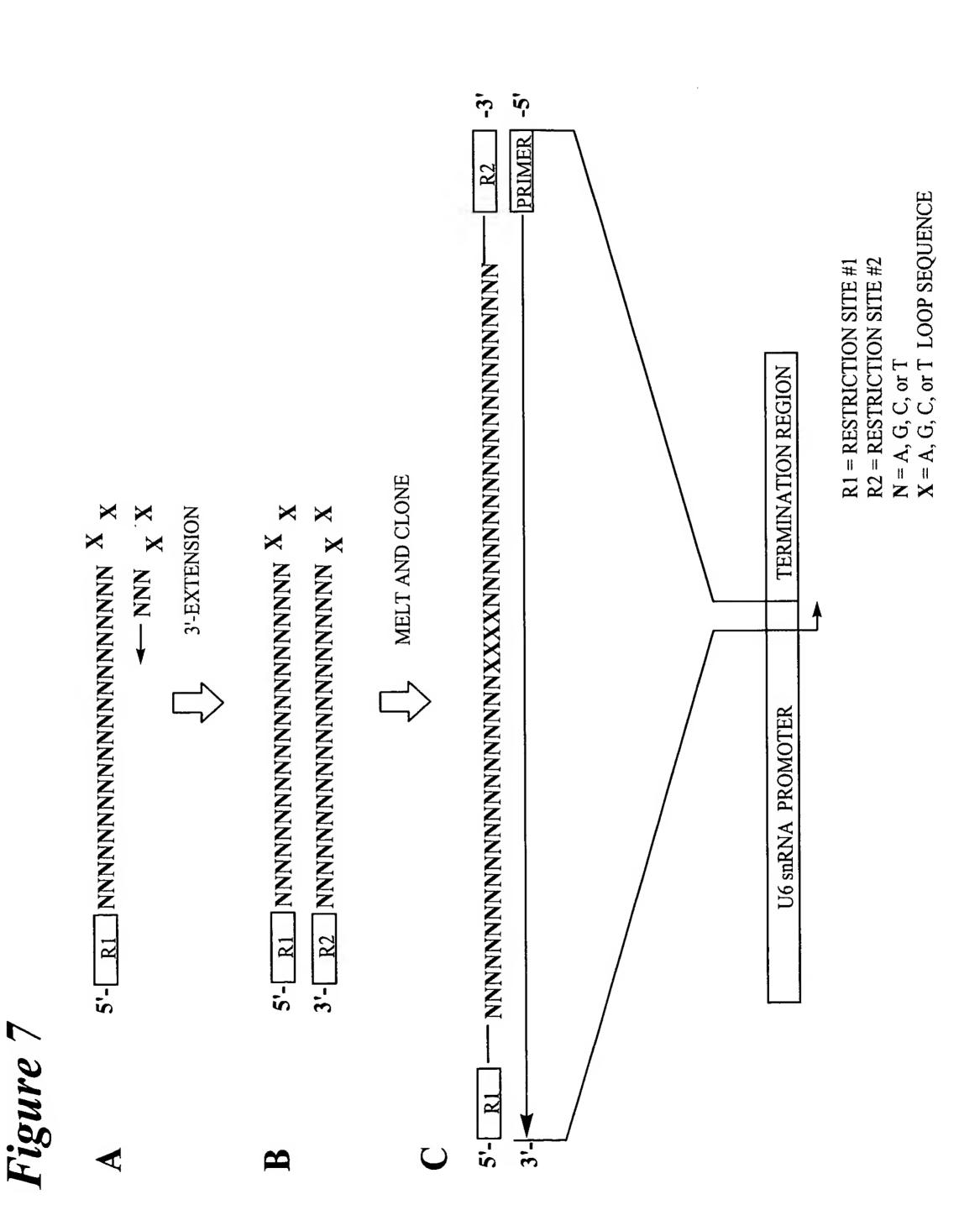


ANTISENSE

SENSE

ANTISENSE

n = 0, 1, 2, 3, 4



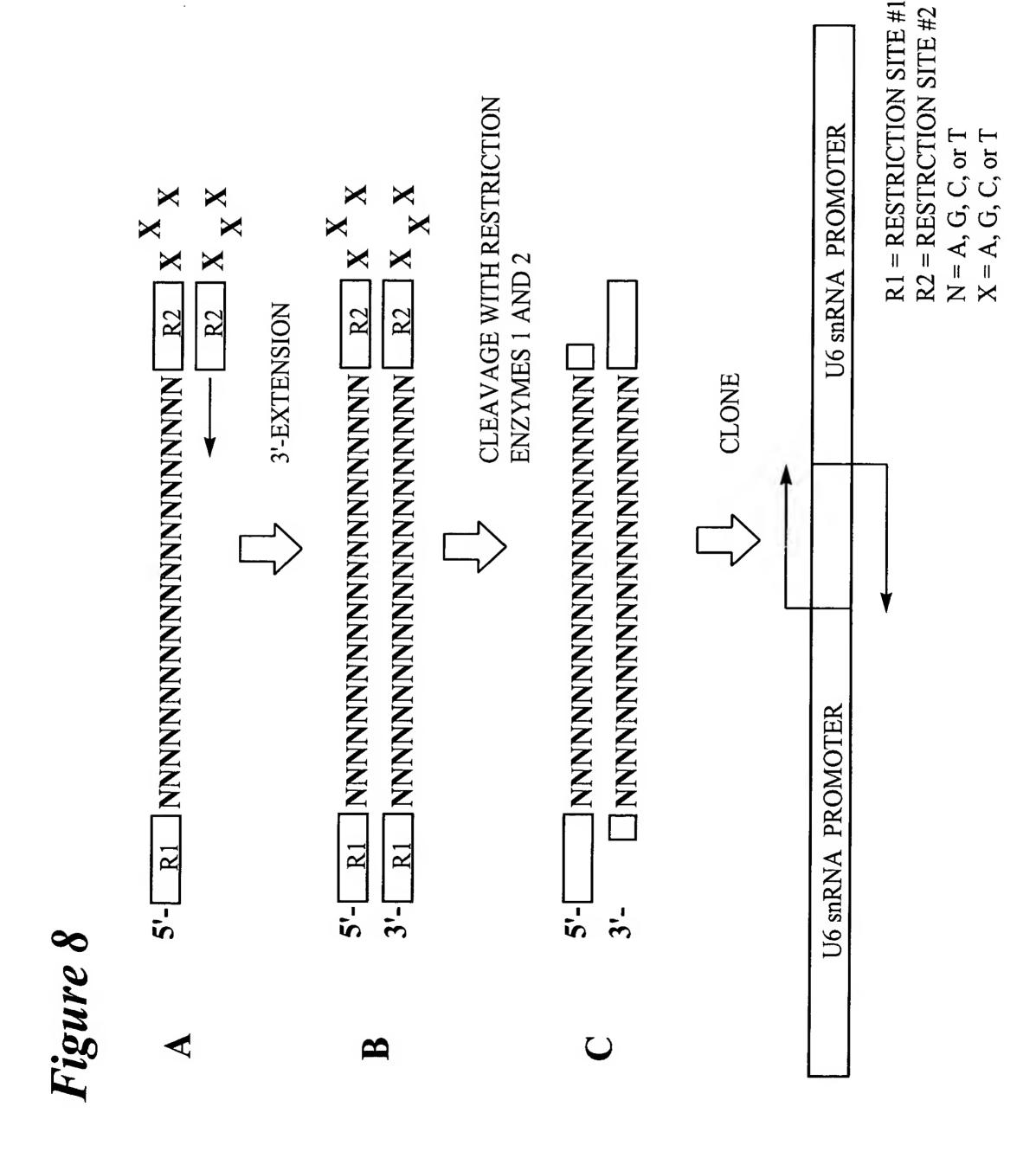
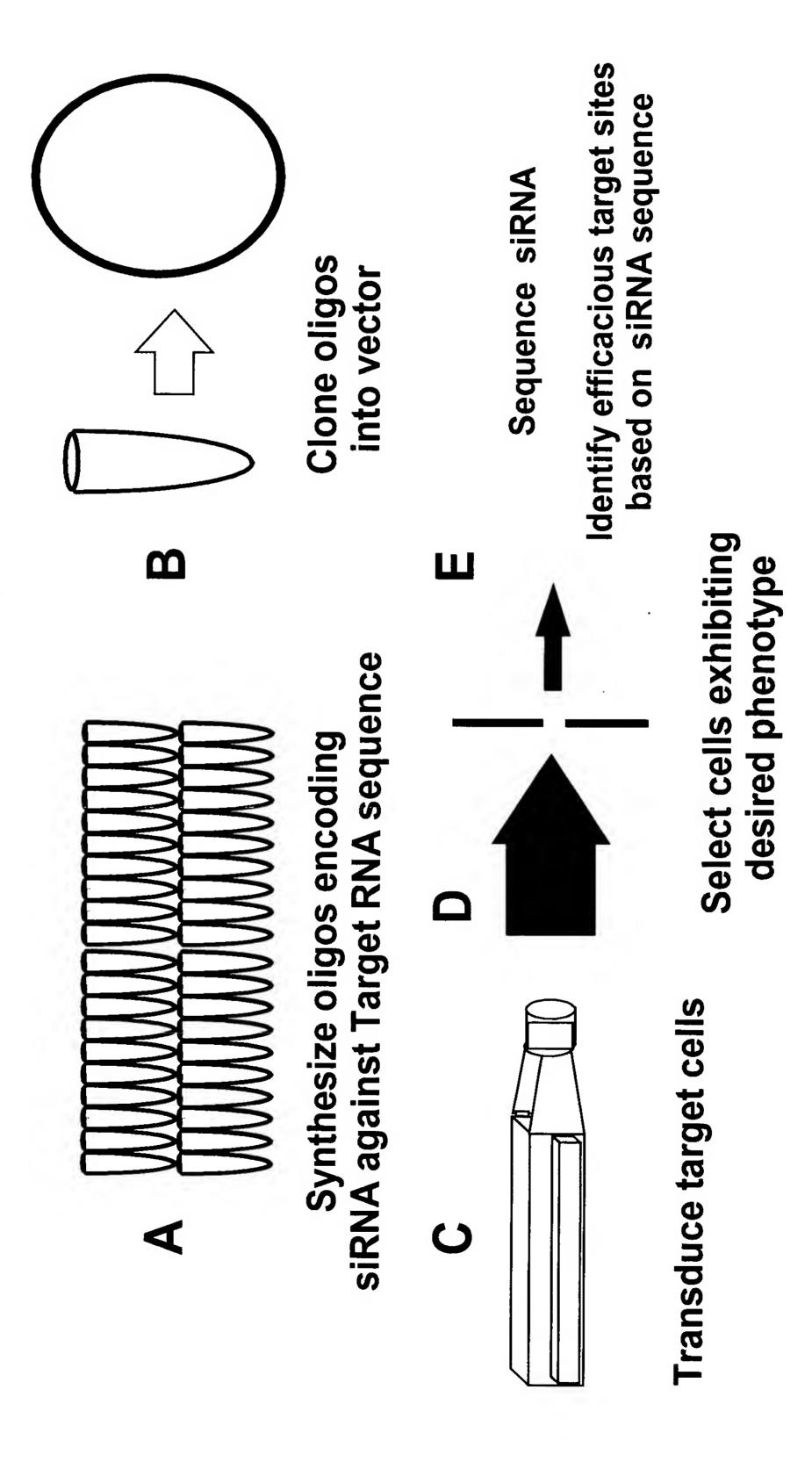
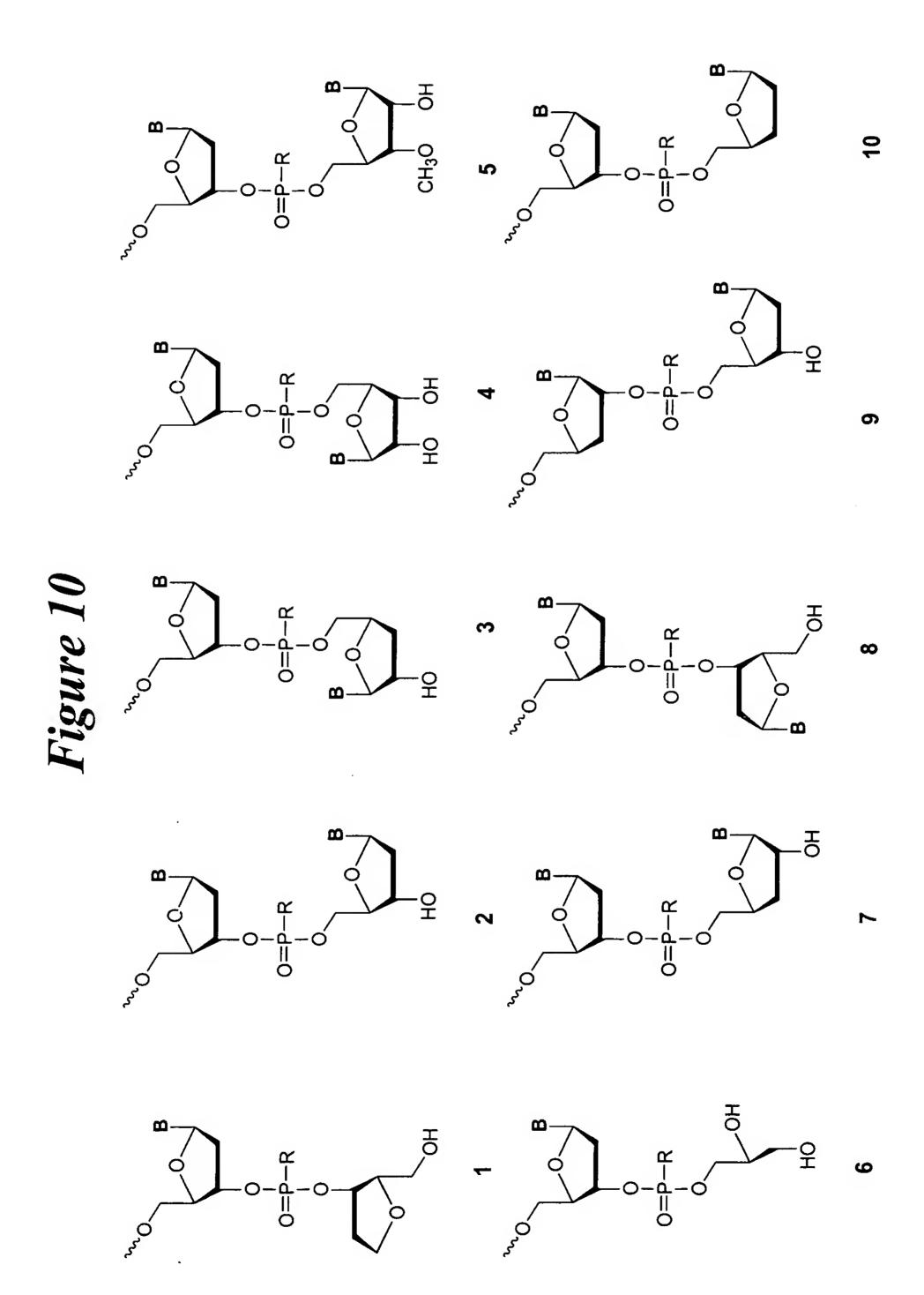


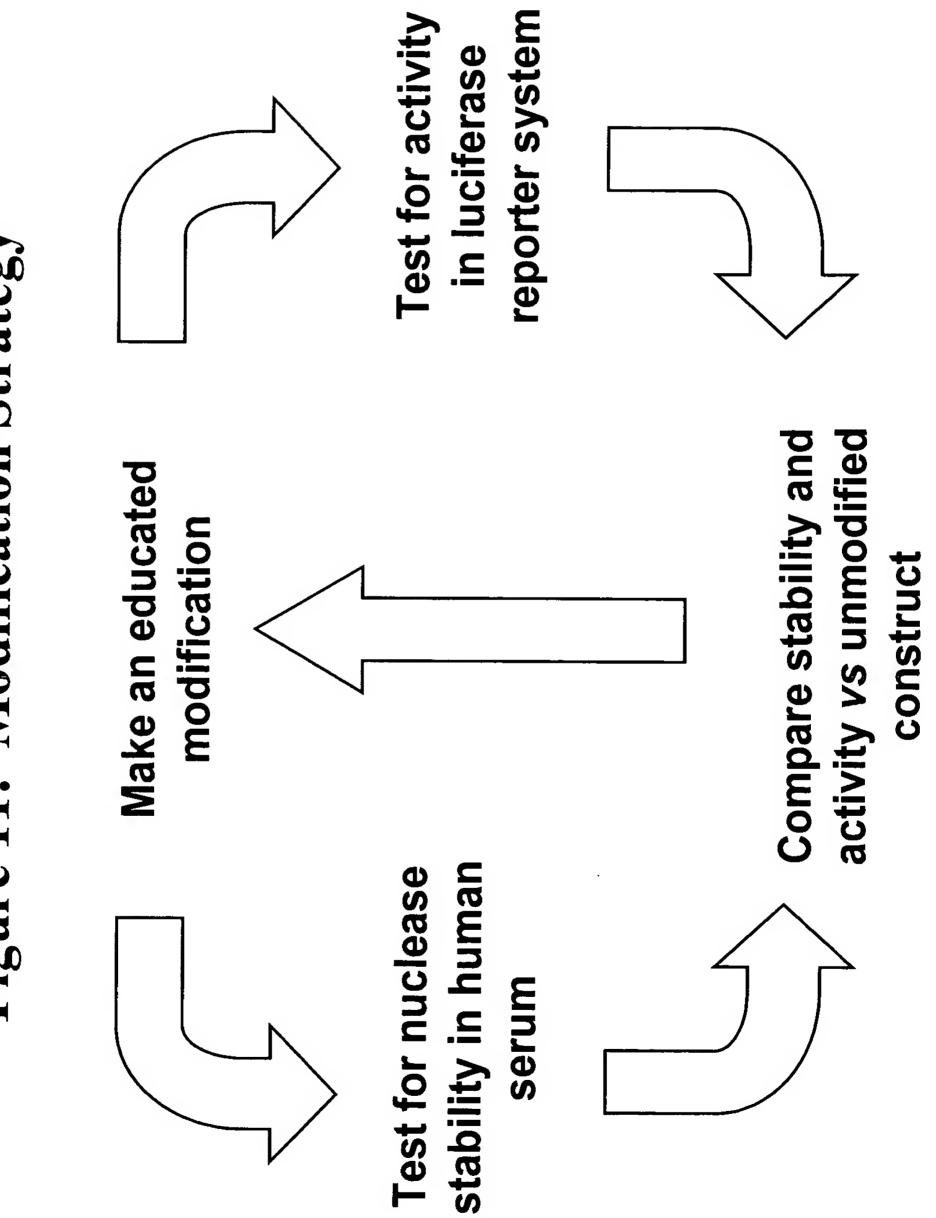
Figure 9: Target site Selection using siRNA





R = O, S, N, alkyl, substituted alkyl, O-alkyl, S-alkyl, alkaryl, or aralkyl B = Independently any nucleotide base, either naturally occurring or chemically modified, or optionally H (abasic).

Figure 11: Modification Strategy



of VEGF-Induced Angiogenesis VEGF Control Inactive 1ug RPI 29983/29984 Inactive 3ug RPI 29983/29984 by siRNAs **p<0.05 with respect to Inverted control by Tukey-Kramer Inactive 10ug RPI 29983/29984 by Dunnett's But svitaA RPI 29695/29699 p< 0.05 with respect to VEGF Inhibition Buc avitaA 요한 29696729699 Active 10ug Figure 12: 요한 29696729699 100 80 9 40 20 0

% Inhibition of VEGF induced sizenegiogenesis

Figure 13: Site 3854 and 3948 KDR RNAi, 4/5, 7/8 and 9/10 chemistry in HAEC cells

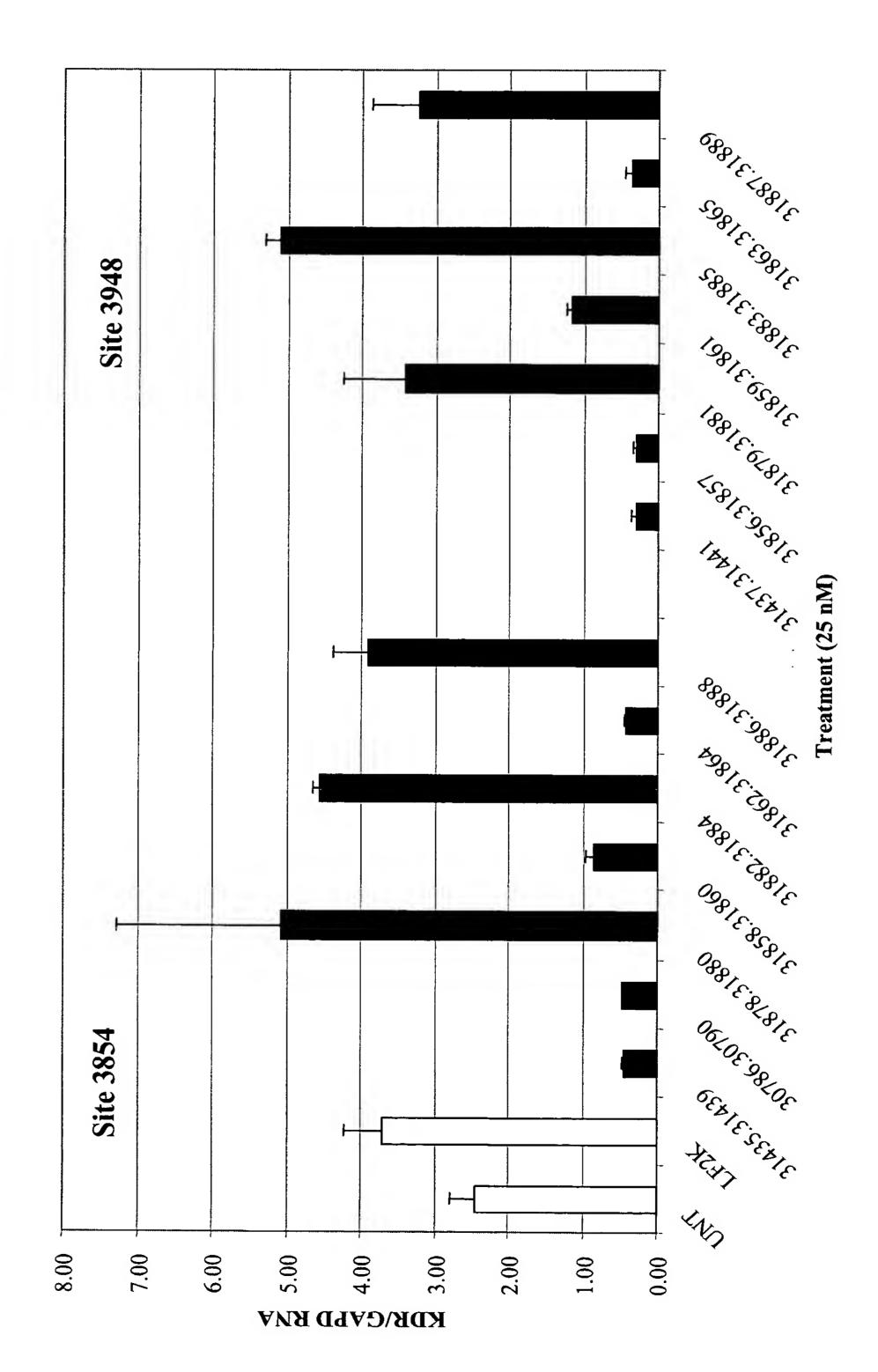
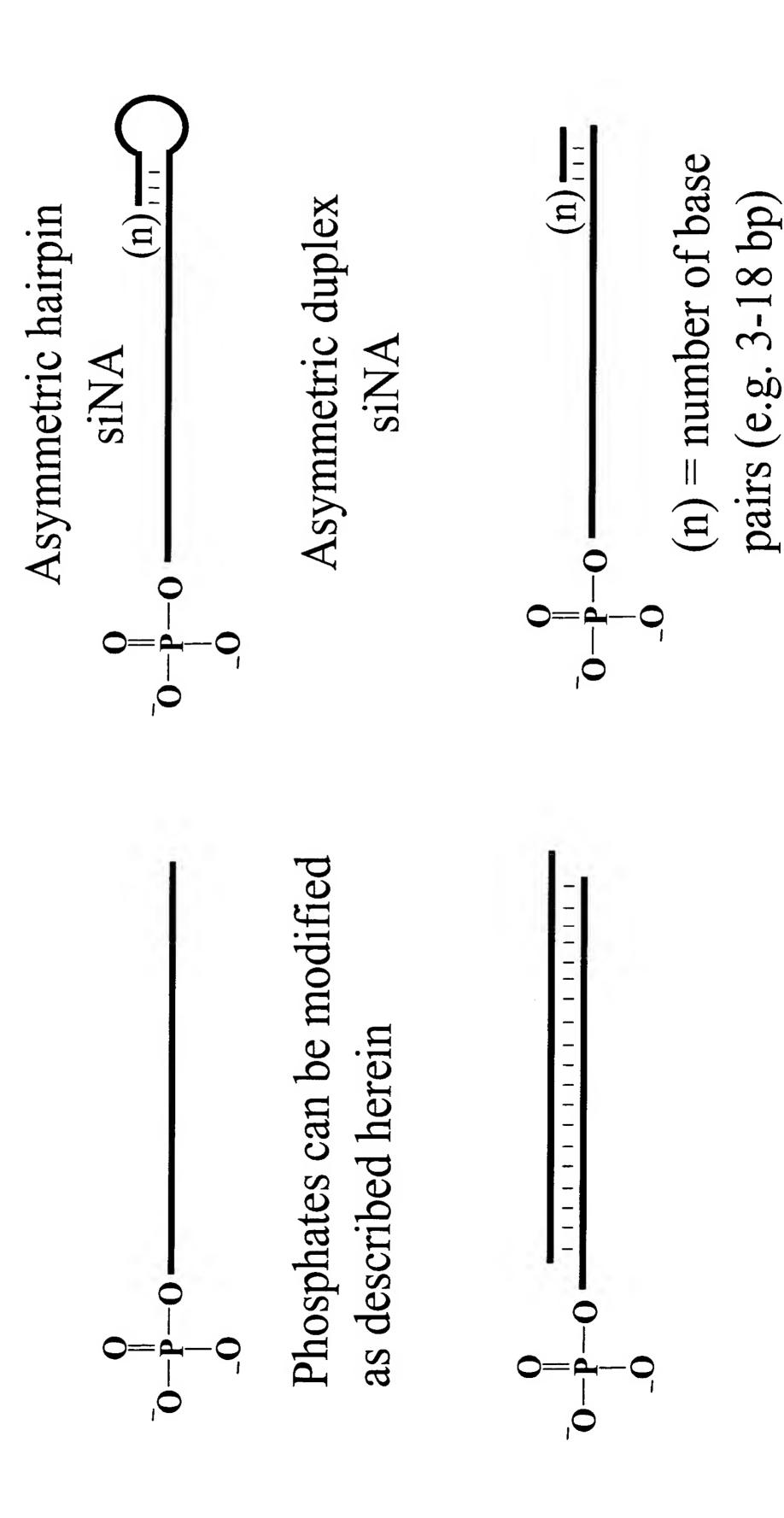


Figure 14: Phosphorylated siNA constructs



combination of other

modifications herein

phosphate modifications Figure 15: 5'

